

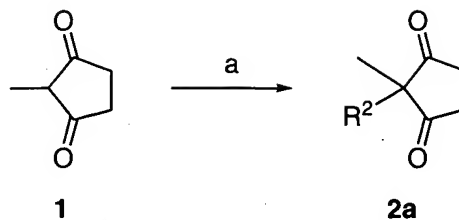


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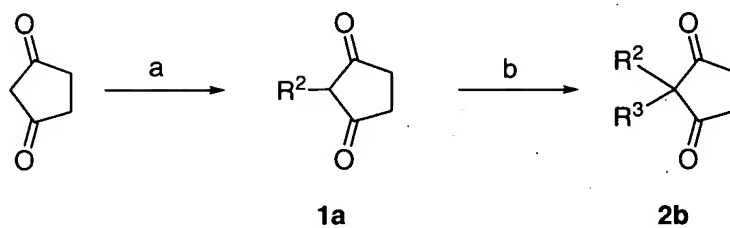
SHEET 1 OF 8 DOCKET 17540CIP (AP)
Serial No. 10/772,720, Applicants: Y. Dondi, et al.
Title: 10,10-DIALKYL PROSTANOIC ACID DERIVATIVES
AS AGENTS FOR LOWERING INTRAOCULAR PRESSURE
714.246.4348 Brent Johnson, ALLERGAN, INC.

Fig. 1

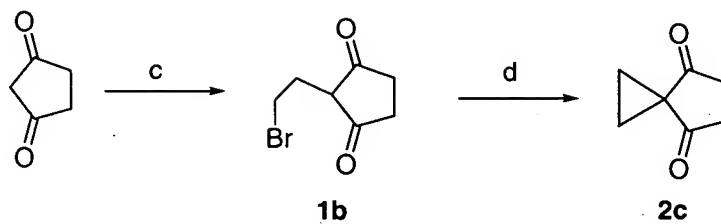
Equation 1



Equation 2



Equation 3

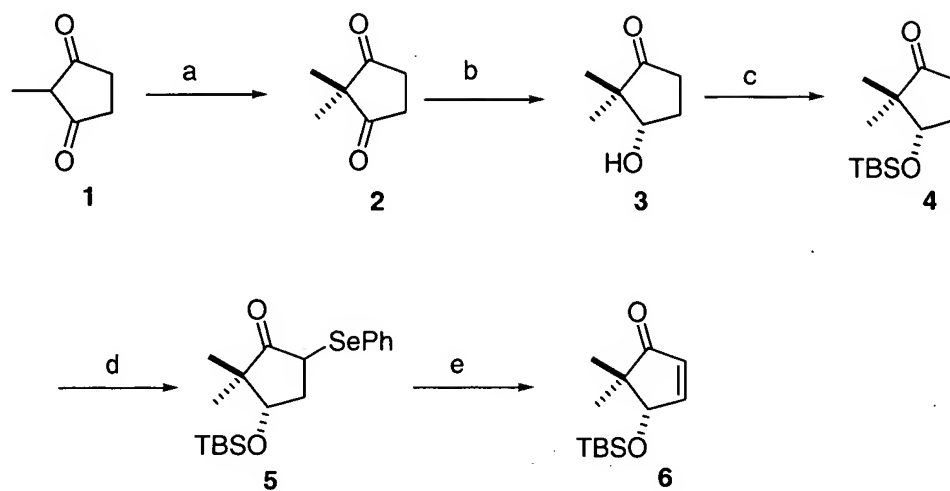


(a) KOH, I-R², dioxane/H₂O; (b) KOH, I-R³, dioxane/H₂O; (c) KOH, 1,2-dibromoethane, dioxane/H₂O; (d) KOH, dioxane/H₂O.

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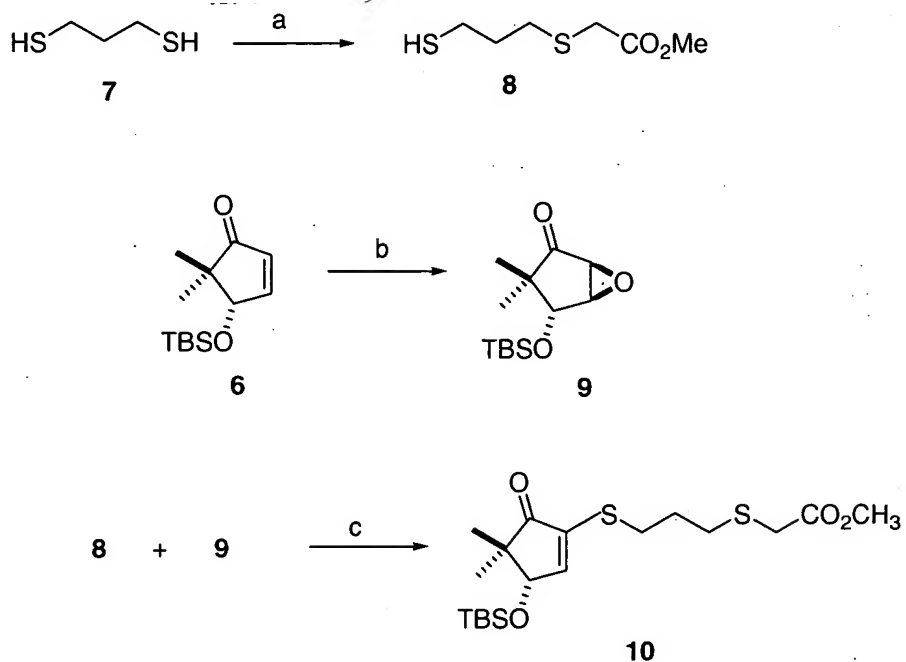
Fig. 2



(a) KOH, MeI, dioxane/H₂O; (b) Baker's Yeast, D-glucose, H₂O; (c) TBSOTf, 2,6-lutidine, CH₂Cl₂; (d) LDA, THF; PhSeCl; (e) 30% H₂O₂, CH₂Cl₂.

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Fig. 3

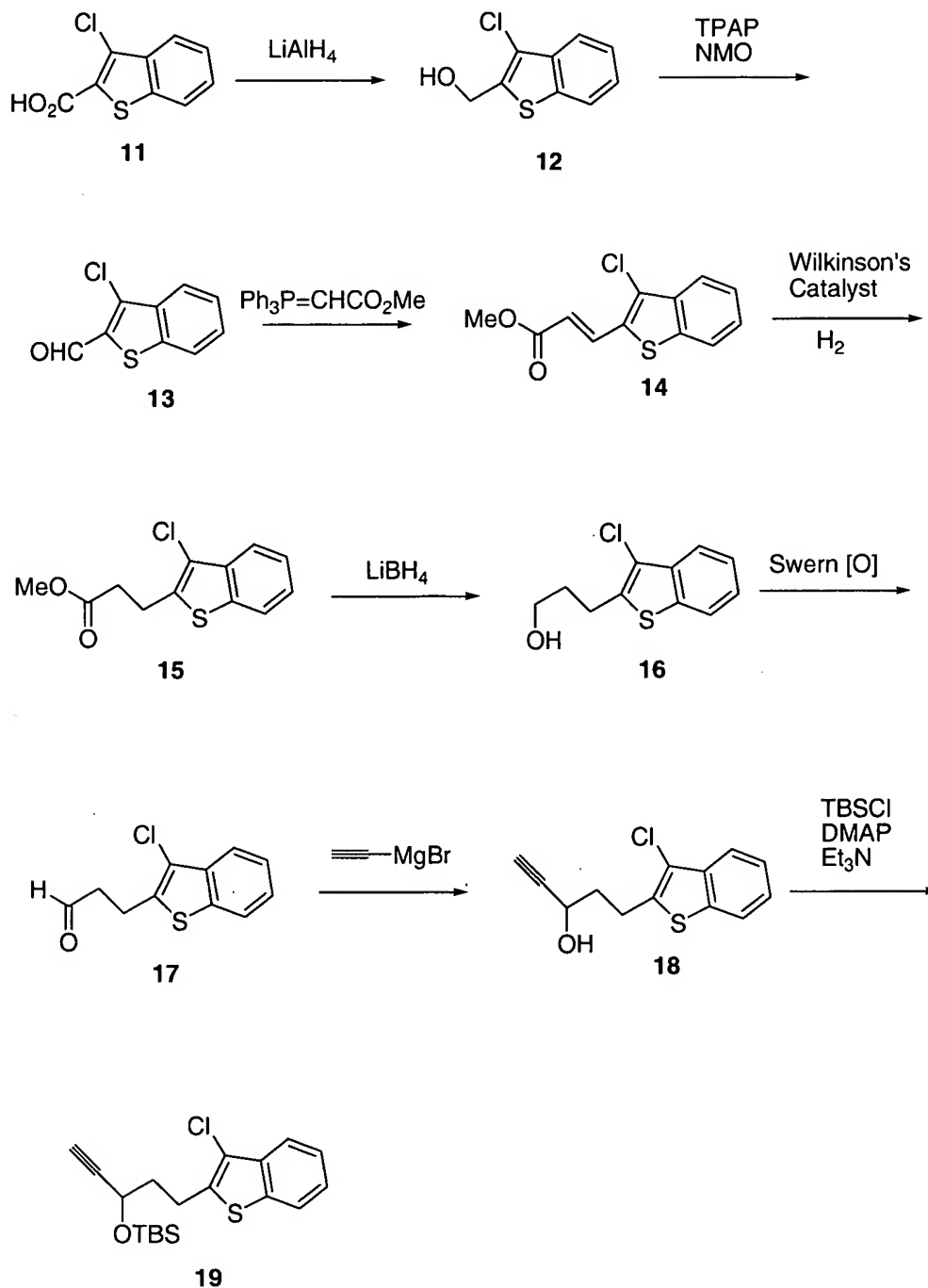


(a) NaH, BrCH₂CO₂CH₃; (b) H₂O₂, NaOH, MeOH; (c) basic Alumina, CH₂Cl₂.

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Fig. 4



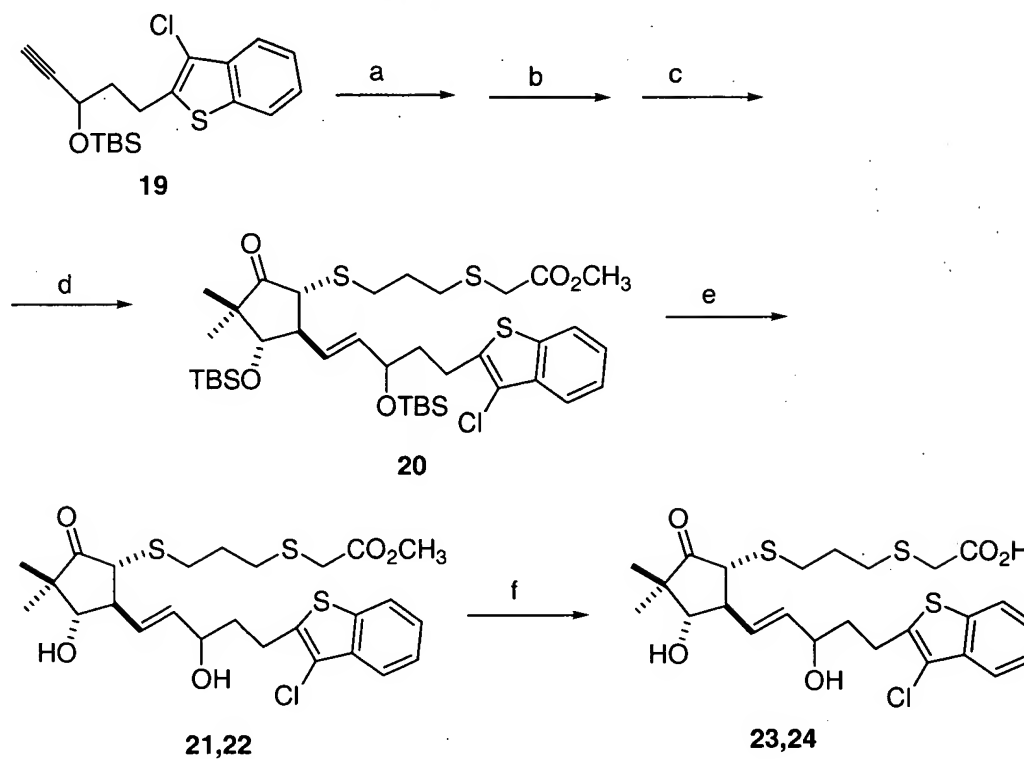
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Fig. 5

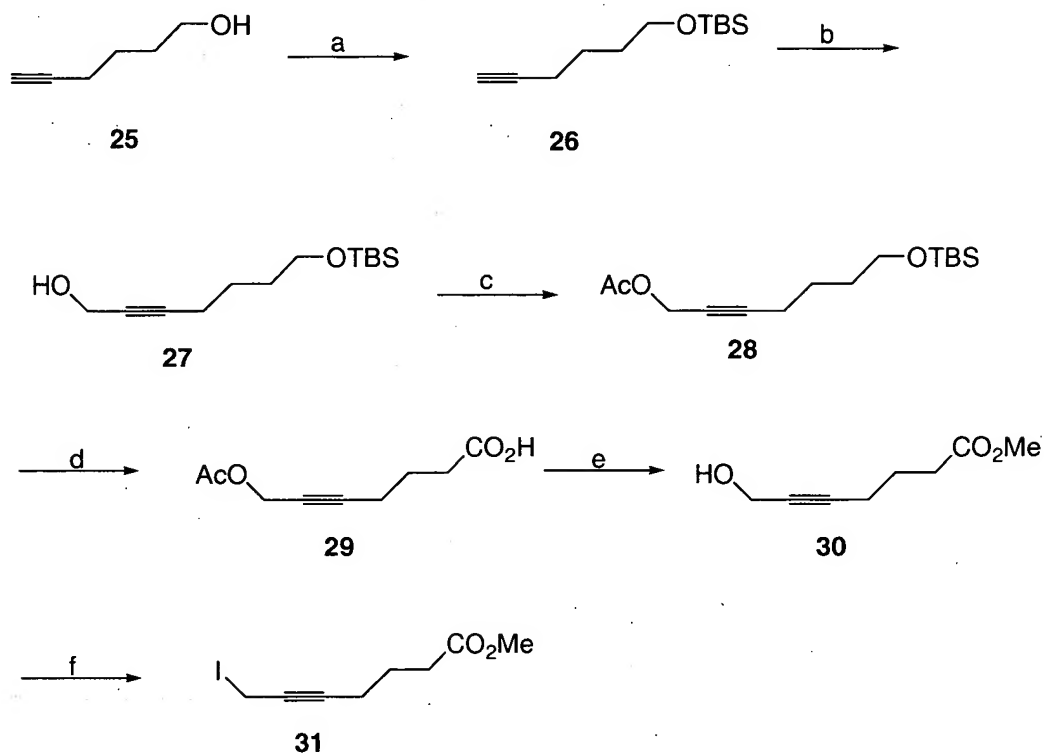


(a) Cp_2ZrHCl , THF; (b) MeLi, Et_2O -78°C ; (c) lithium 2-thienycyanocuprate;
(d) enone 10, THF -78°C ; (e) HF-pyridine, CH_3CN ; separate diastereomers
(f) rabbit liver esterase, phosphate buffer, CH_3CN .

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Fig. 6

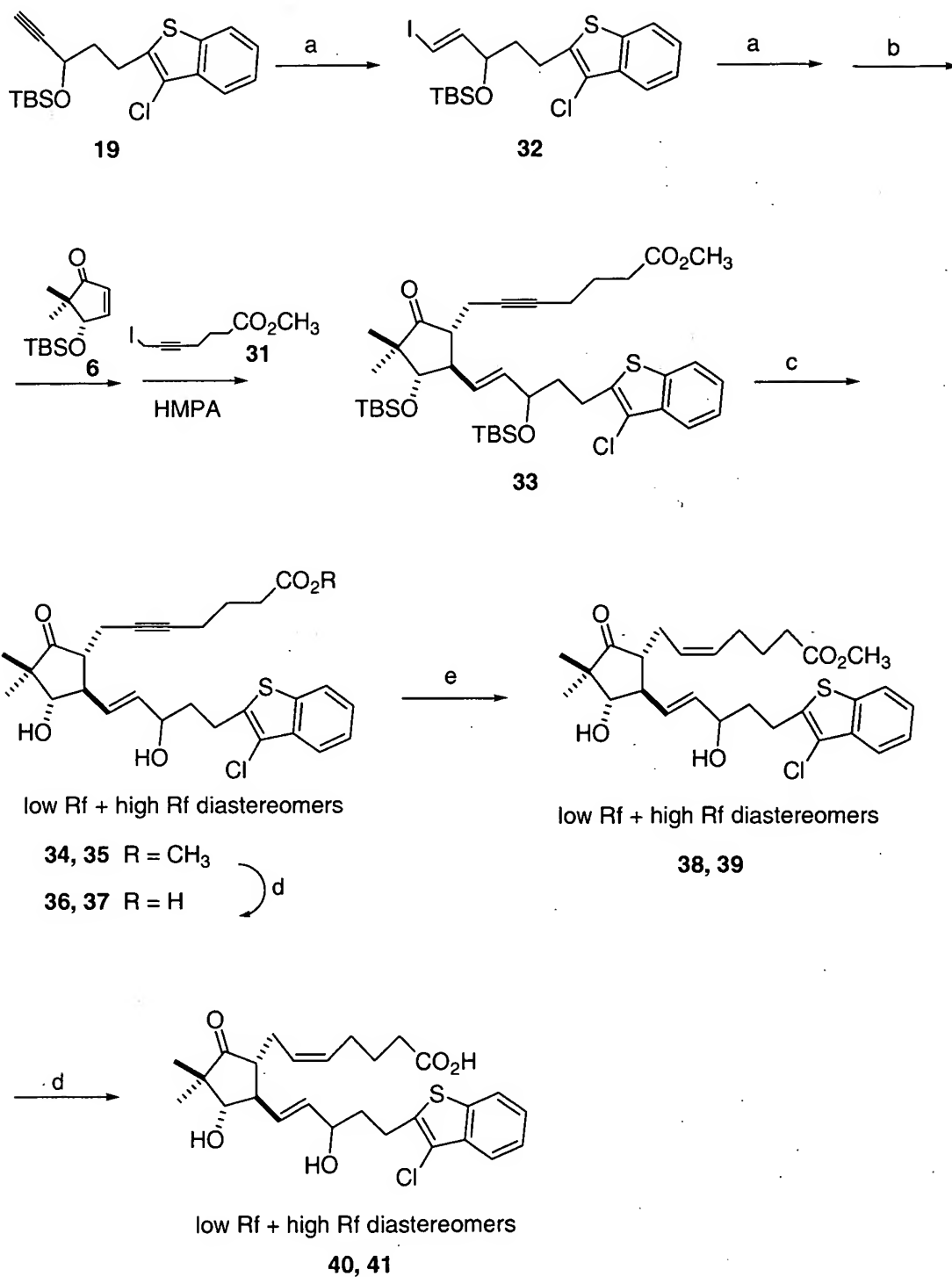


(a) TBSCl, etc.; (b) *n*-BuLi; DMF; (c) Ac₂O, pyridine; (d) Jones oxidation; (e) MeOH, AcCl; (f) PPh₃, I₂, imidazole, CH₂Cl₂.

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Fig. 7

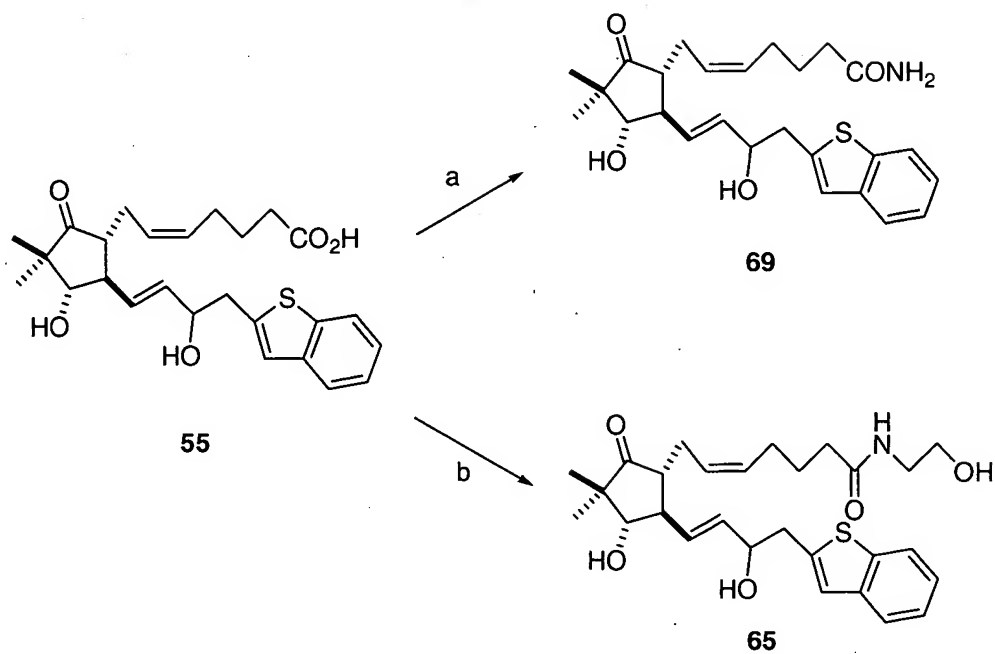


(a) *t*-BuLi, THF -78 °C; (b) Me₂Zn; (c) HF-pyridine, CH₃CN; separate diastereomers; (d) rabbit liver esterase, pH 7.2 phosphate buffer, CH₃CN; (e) NiCl₂, NaBH₄, ethylenediamine, H₂, THF;

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Fig. 8



(a) $\text{ClCO}_2\text{CH}_2\text{CH}_3$, Et_3N , CH_2Cl_2 ; NH_4OH (aq);
 (b) EDCI , N -hydroxysuccinimide, $\text{H}_2\text{NCH}_2\text{CH}_2\text{OH}$, DMF .